Pippig et al.
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IN THE CLAIMS

As set forth below, please cancel claims 1-8, 10-19, 21-22, 28, and 30-35; amend claims 20, 23-27, 29, 36, and 39; and add new claims 43-72:

1-19. (Cancelled)

20. (Currently amended) A method for identifying transduced mammalian <u>hematopoietic</u> cells comprising:



- a) retrovirally transducing mammalian hematopoietic cells with a nucleic acid sequence encoding a protein-tyrosine-kinase-receptor (PTKR) mutated epidermal growth factor receptor (EGFR) operatively linked to an expression control sequence, wherein said mutated EGFR PTKR comprises modifications to the intracellular and the extracellular domains, comprises a modification to the extracellular domain, or excludes any nerve growth factor-receptor(s) (NGFR) and comprises a modification to the intracellular domain;
- b) incubating the transduced mammalian <u>hematopoietic</u> cells with a marked antibody which recognizes and binds specifically to the mutated EGFR PTKR; and
- c) identifying the marked transduced mammalian hematopoietic cells.

21-22. (Cancelled)

- 23. (Currently amended) The method according to claim 20, wherein the mammalian hematopoietic cells are transduced by a retroviral vector selected from the group consisting of a moloney murine leukemia viral vector, a myeloproliferative sarcoma viral vector, a murine embryonic stem cell viral vector, a murine stem cell viral vector, and a spleen focus forming viral vector.
- 24. (Currently amended) The method according to claim 20, wherein the mammalian hematopoietic cells are transduced by a lentiviral vector.

- 25. (Currently amended) The method according to claim 20, further comprising the step of separating the identified marked transduced mammalian <u>hematopoietic</u> cells from non-marked mammalian <u>hematopoietic</u> cells.
- 26. (Currently amended) The method according to claim 20, further comprising the step of expanding the marked transduced mammalian <u>hematopoietic</u> cells.
- 27. (Currently amended) A method of identifying mammalian <u>hematopoietic</u> cells expressing a protein of interest, comprising:
 - a) introducing into a mammalian hematopoietic cell a nucleic acid comprising a DNA sequence encoding a protein of interest and comprising a DNA sequence encoding a protein-tyrosine kinase receptor (PTKR), mutated epidermal growth factor receptor (EGFR), wherein said DNA sequences are operatively linked to one or more expression control sequences, and wherein said mutated EGFR PTKR comprises modifications to the intracellular and the extracellular domains, comprises a modification to the extracellular domain, or excludes any nerve growth factor receptor(s) (NGFR) and comprises a modification to the intracellular domain;
 - b) culturing the resulting mammalian hematopoietic cells; and
 - c) identifying mammalian <u>hematopoietic</u> cells which express the mutated EGFR PTKR
 thereby obtaining mammalian <u>hematopoietic</u> cells which express the protein of
 interest.
- 28. (Cancelled)
- (Currently amended) The method according to claim 27, wherein the nucleic acid encoding the mutated PTKR and the nucleic acid encoding the protein of interest are is introduced by a retroviral vector.

30-35. (Cancelled)

36. (Currently amended) The method according to claim <u>20</u> 22, wherein the mutated EGFR is a mutated EGFR1.

- 37. (Previously added) The method according to claim 36, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 679-1210 are deleted.
- 38. (Previously added) The method according to claim 36, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 25-312 and 679-1210 are deleted.



- 39. (Currently amended) The method according to claim <u>27</u> 28, wherein the mutated EGFR is a mutated EGFR 1.
- 40. (Previously added) The method according to claim 39, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 679-1210 are deleted.
- 41. (Previously added) The method according to claim 39, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 25-312 and 679-1210 are deleted.
- 42. (Previously added) A method of identifying a genetically modified mammalian cell, comprising:
 - a) introducing a nucleic acid sequence encoding a mutated epidermal growth factor receptor 1 (EGFR1), operatively linked to an expression control sequence, into a mammalian cell to form a genetically modified mammalian cell, wherein the mutated EGFR1 either comprises: i) the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 679-1210 are deleted, or ii) the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 25-312 and 679-1210 are deleted;
 - b) allowing expression of the mutated EGFR1 in the genetically modified mammalian cell; and
 - c) identifying said genetically modified mammalian cell expressing the mutated EGFR1.

- 43. (New) The method according to claim 42, wherein the introducing step is accomplished by incorporating the nucleic acid sequence encoding the mutated EGFR1 into a vector and introducing said vector into said mammalian cell.
- 44. (New) The method according to claim 43, wherein the vector is a retroviral vector.

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- 45. (New) The method according to claim 42, further comprising separating the identified cell expressing the mutated EGFR1.
- 46. (New) The method according to claim 42, wherein the mammalian cell is a human cell.
- 47. (New) The method according to claim 46, wherein the human cell is selected from the group consisting of a hematopoietic cell, a liver cell, an endothelial cell and a smooth muscle cell.
- 48. (New) The method according to claim 46, wherein the human cell is a hematopoietic cell.
- 49. (New) The method according to claim 48, wherein the hematopoietic cell is a stem cell or T-cell.
- 50. (New) The method according to claim 43, wherein a heterologous coding sequence is also incorporated into said vector.
- 51. (New) The method according to claim 42, wherein the identifying step is accomplished by contacting the genetically modified mammalian cell with an antibody that recognizes and binds to the mutated EGFR1.

- 52. (Re-presented formerly claim 12) A method of identifying a genetically modified cell, comprising:
 - a) introducing into a cell a nucleic acid sequence encoding a mutated epidermal growth factor receptor (EGFR), operatively linked to an expression control sequence, to form a genetically modified cell, wherein said cell is a human hematopoietic cell, a human liver cell, a human endothelial cell, or a human smooth muscle cell, and wherein said mutated EGFR comprises modifications to the intracellular and the extracellular domains, comprises a modification to the extracellular domain, or comprises a modification to the intracellular domain;
 - b) allowing expression of the mutated EGFR in the genetically modified cell; and
 - c) identifying said genetically modified cell expressing the mutated EGFR.
- 53. (New) The method according to claim 52, wherein the cell is a human hematopoietic cell.
- 54. (New) The method according to claim 53, wherein the human hematopoietic cell is a stem cell or T-cell.
- 55. (New) The method according to claim 52, wherein the mutated EGFR is a mutated EGFR1.
- 56. (New) The method according to claim 55, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 679-1210 are deleted.
- 57. (New) The method according to claim 55, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 25-312 and 679-1210 are deleted.
- 58. (New) The method according to claim 52, wherein the introducing step is accomplished by incorporating the nucleic acid sequence encoding the mutated EGFR into a vector and introducing said vector into said cell.



- 59. (New) The method according to claim 58, wherein the vector is a retroviral vector.
- 60. (New) The method according to claim 52, further comprising separating the identified cell expressing the mutated EGFR.
- 61. (New) The method according to claim 58, wherein a heterologous coding sequence is also incorporated into said vector.
- 62. (New) The method according to claim 52, wherein the identifying step is accomplished by contacting the genetically modified cell with an antibody that recognizes and binds to the mutated EGFR.
- 63. (Re-presented formerly dependent claim 24) A method for identifying transduced mammalian cells comprising:
 - a) transducing mammalian cells with a lentiviral vector comprising a nucleic acid sequence encoding a mutated epidermal growth factor receptor (EGFR) operatively linked to an expression control sequence, wherein said mutated EGFR comprises modifications to the intracellular and the extracellular domains, comprises a modification to the extracellular domain, or comprises a modification to the intracellular domain;
 - b) incubating the transduced mammalian cells with a marked antibody which recognizes and binds specifically to the mutated EGFR; and
 - c) identifying the marked transduced mammalian cells.
- 64. (New) The method according to claim 63, wherein the mutated EGFR is a mutated EGFR1.
- 65. (New) The method according to claim 64, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 679-1210 are deleted.
- 66. (New) The method according to claim 64, wherein the mutated EGFR1 comprises the amino acid sequence set forth in SEQ ID NO:2 except that amino acid residues 25-312 and 679-1210 are deleted.

- 67. (New) The method according to claim 63, wherein the mammalian cells are human cells.
- 68. (New) The method according to claim 67, wherein the human cells are selected from the group consisting of hematopoietic cells, liver cells, endothelial cells and smooth muscle cells.
- 69. (New) The method according to claim 67, wherein the human cells are hematopoietic cells.
- 70. (New) The method according to claim 69, wherein the hematopoietic cells are stem cells or T-cells.
- 71. (New) The method according to claim 63, wherein said lentiviral vector also comprises a nucleic acid sequence encoding a heterologous protein of interest.
- 72. (New) The method according to claim 63, wherein the identifying step is accomplished by contacting the genetically modified mammalian cell with an antibody that recognizes and binds to the mutated EGFR.